



FORM PTO-1449 (REV. 7-80)	U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE	ATTY. DOCKET NO.	APPLICATION NO.
		720109.401	10/669,158
		APPLICANTS	
		Edward A. Neuwalt et al.	
INFORMATION DISCLOSURE STATEMENT (Use several sheets if necessary)		FILING DATE	GROUP ART UNIT
		September 22, 2003	1614

U.S. PATENT DOCUMENTS

*EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
EA	AA	5,002,935	03/26/91	Bodor	514	58	
EA	AB	5,601,835	02/11/97	Sabel et al.	424	424	
EA	AC	5,648,331	07/15/97	Koudsi et al.	514	12	
EA	AD	5,869,044	02/09/99	Tomaru et al.	424	74.64	
EA	AE	5,912,019	06/15/99	Singh	424	608	
EA	AF	5,968,959	10/19/99	Haikala et al.	514	345	
EA	AG	6,001,842	12/14/99	Neely	514	263	
EA	AH	6,086,868	07/11/00	Moldawer et al.	424	85.2	
	AI						

FOREIGN PATENT DOCUMENTS

		DOCUMENT NUMBER	DATE	COUNTRY	TRANSLATION	
					YES	NO
	AJ					
	AK					

OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

EA	AL	Andersen, L.W. et al., "The role of N-acetylcystein administration on the oxidative response of neutrophils during cardiopulmonary bypass," <i>Perfusion</i> 10: 21-26, 1995.
EA	AM	Ataka, K. et al., "The Protective Effects of Calcium Antagonist and Free Radical Scavenger Against Myocardial Ischemic/Reperfusion Injury in the Isolated Rat Heart," <i>Kobe J. Med. Sci.</i> 35(5-6): 261-276, December 1989.
EA	AN	Auer, M. et al., "Effects of Clinically Used Antioxidants in Experimental Pneumococcal Meningitis," <i>The Journal of Infectious Diseases</i> 182: 347-350, July 2000.
EA	AO	Barbut, D. et al., "Aortic Atheroma Is Related to Outcome but Not Numbers of Emboli During Coronary Bypass," <i>Ann. Thorac. Surg.</i> 64: 454-459, 1997.
EA	AP	Blakley, B.W. et al., "Strategies for Prevention of Toxicity Caused by Platinum-Based Chemotherapy: Review and Summary of the Annual Meeting of the Blood-Brain Barrier Disruption Program, Gleneden Beach, Oregon, March 10, 2001," <i>The Laryngoscope</i> 112(11): 1997-2001, November 2002.

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	BA					
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	BB					
OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)						
EA	BC	Börjesson, A. et al., "Effects of N-Acetylcysteine on Pulmonary Macrophage Activity after Intestinal Ischemia and Reperfusion in Rats," <i>Digestive Surgery</i> 17: 379-389, 2000.				
EA	BD	Brunet, J. et al., "Effects of N-Acetylcysteine in the Rat Heart Reperfused After Low-flow Ischemia: evidence for a Direct Scavenging of Hydroxyl Radicals and a Nitric Oxide-Dependent Increase in Coronary Flow," <i>Free Radical Biology & Medicine</i> 19(5): 627-638, 1995.				
EA	BE	Cakir, O. et al., "Neuroprotective effect of N-acetylcysteine and hypothermia on the spinal cord ischemia-reperfusion injury," <i>Cardiovascular Surgery</i> 11(5): 375-379, October 2003.				
EA	BF	Cao and Phillis, "The free radical scavenger, α -lipoic acid, protects against cerebral ischemia-reperfusion injury in gerbils," <i>Free Rad. Res.</i> 23(4): 365-370, 1995.				
EA	BG	Carroll, J.E. et al., "Nuclear factor- κ B activation during cerebral reperfusion: effect of attenuation with N-acetylcysteine treatment," <i>Molecular Brain Research</i> 56: 186-191, 1998.				
EA	BH	Ceconi, C. et al., "The Role of Glutathione Status in the Protection Against Ischaemic and Reperfusion Damage: Effects of N-Acetyl Cysteine," <i>J. Mol. Cell. Cardiol.</i> 20: 5-13, 1988.				
EA	BI	Chavez-Cartaya, R. et al., "Free radical scavengers to prevent reperfusion injury following experimental warm liver ischaemia. Is there a real physiological benefit?" <i>Transplant International</i> 12(3): 213-221, 1999.				
EA	BJ	Christopher, T.A. et al., "Carvedilol, a New Beta-Adrenoreceptor Blocker, Vasodilator and Free-Radical Scavenger, Exerts an Anti-Shock and Endothelial Protective Effect in Rat Splanchnic Ischemia and Reperfusion," <i>The Journal of Pharmacology and Experimental Therapeutics</i> 273(1): 64-71, April 1995.				
EA	BK	Cosgrove, D.M. et al., "Primary myocardial revascularization. Trends in surgical mortality," <i>The Journal of Thoracic and Cardiovascular Surgery</i> 88(5 pt. 1): 673-684, November 1984.				
EA	BL	Cuzzocrea, S. et al., "Beneficial effects of n-acetylcysteine on ischaemic brain injury," <i>British Journal of Pharmacology</i> 130: 1219-1226, 2000.				
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	CA						
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		DOCUMENT NUMBER	DATE	COUNTRY		TRANSLATION YES NO	
	CB						
OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)							
EA	CC	Cuzzocrea, S. et al., "Effects on n-acetylcysteine in a rat model of ischemia and reperfusion injury," <i>Cardiovascular Research</i> 47: 537-548, 2000.					
EA	CD	Dahlborg et al., "The Potential for Complete and Durable Response in Nonglial Primary Brain Tumors in Children and Young Adults with Enhanced Chemotherapy Delivery," <i>The Cancer Journal from Scientific American</i> 4(2): 110-124, March/April 1998.					
EA	CE	Demir, S. et al., "Pentoxifylline and N-acetylcysteine in hepatic ischemia/reperfusion injury," <i>Clinica Chimica Acta</i> 275: 127-135, 1998.					
EA	CF	DiMari, J. et al., "N-acetyl cysteine ameliorates ischemic renal failure," <i>American Journal of Physiology</i> 272(3 pt. 2): F292-F298, March 1997.					
EA	CG	Doolittle, N.D. et al., "Delayed Sodium Thiosulfate as an Otoprotectant Against Carboplatin-induced Hearing Loss in Patients with Malignant Brain Tumors," <i>Clinical Cancer Research</i> 7: 493-500, March 2001.					
EA	CH	Ellis, E.F. et al., "Restoration of cerebrovascular responsiveness to hyperventilation by the oxygen radical scavenger n-acetylcystein following experimental traumatic brain injury," <i>Journal of Neurosurgery</i> 75(5): 774-779, November 1991.					
EA	CI	Fukahara, K. et al., "Inhaled nitric oxide after lung ischemia reperfusion; effect on hemodynamics and oxygen free radical scavenger system," <i>European Journal of Cardio-thoracic Surgery</i> 11:343-349, 1997.					
EA	CJ	Fukuzawa, K. et al., "N-Acetylcystein Ameliorates Reperfusion Injury After Warm Hepatic Ischemia," <i>Transplantation</i> 59(1): 6-9, January 15, 1995.					
EA	CK	Grieb, P. et al., "Changes in oxidative stress in the rat brain during post-cardiac arrest reperfusion, and the effect of treatment with the free radical scavenger idebenome," <i>Resuscitation</i> 39: 107-113, 1998.					
EA	CL	Harrison, P.M. et al., "Improvement by acetylcysteine of hemodynamics and oxygen transport in fluminant hepatic failure," <i>The New England Journal of Medicine</i> 324(26): 1852-1857, June 27, 1991.					
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				YES	NO	
DB						
OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)						
EA	DC	Hori, K. et al., "Neuroprotection by glial cells through adult T cell leukemia-derived factor/human thioredoxin (ADF/TRX)," <i>Brain Research</i> 652(2): 304-310, August 1994.				
EA	DD	Horowitz, J.D., "Thiol-Containing Agents in the Management of Unstable Angina Pectoris and Acute Myocardial Infarction," <i>The American Journal of Medicine</i> 91(3C): 113s-117s, September 30, 1991.				
EA	DE	Hur, G.M. et al., "Hepatic Ischemia/Reperfusion in Rats Induces iNOS Gene Transcription by Activation of NF- κ B," <i>Biochemical and Biophysical Research Communications</i> 261(3): 917-922, 1999.				
EA	DF	Jolliet-Riant and Tillement, "Drug transfer across the blood-brain barrier and improvement of brain delivery," <i>Fundam. Clin. Pharmacol.</i> 13: 16-26, 1999.				
EA	DG	Kannan, R. et al., "Transport of Glutathione at Blood-Brain Barrier of the Rat: Inhibition by Glutathione Analogs and Age-Dependence," <i>Journal of Pharmacology and Experimental Therapeutics</i> 263(3): 964-970, December 1992.				
EA	DH	Knuckey, N.W. et al., "N-Acetylcysteine Enhances Hippocampal Neuronal Survival After Transient Forebrain Ischemia in Rats," <i>Stroke</i> 26: 305-311, 1995.				
EA	DI	Koeppel, T.A. et al., "Impact of N-Acetylcysteine on the Hepatic Microcirculation After Orthotopic Liver Transplantation," <i>Transplantation</i> 61(9): 1397-1402, May 15, 1996.				
EA	DJ	Koeppel, T.A. et al., "Improvement of Hepatic Microhemodynamics by N-Acetylcysteine after Warm Ischemia," <i>European Surgical Research</i> 28: 270-277, 1996.				
EA	DK	Koksal, C. et al., "Attenuation of Ischemia/Reperfusion Injury by N-Acetylcysteine in a Rat Hind Limb Model," <i>Journal of Surgical Research</i> 111(2): 236-239, May 15, 2003.				
EA	DL	Kretzschmar, M. et al., "Reduction of ischemia-reperfusion syndrome after abdominal aortic aneurysmectomy by N-acetylcysteine but not mannitol," <i>Acta Anaesthesiologica Scandinavica</i> 40(6): 657-664, July 1996.				
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EA						

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				YES	NO
EB					

OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

EA	EC	Kreuter, J., "Nanoparticulate systems for brain delivery of drugs," <i>Advanced Drug Delivery Reviews</i> 47: 65-81, 2001.
EA	ED	Kuluz, J.W. et al., "The Effect of Nitric Oxide Synthase Inhibition on Infarct Volume After Reversible Focal Cerebral Ischemia in Conscious Rats," <i>Stroke</i> 24(12): 2023-2029, December 1993.
EA	EE	Lyrer, P. et al., "Levels of low molecular weight scavengers in the rat brain during focal ischemia," <i>Brain Research</i> 567(2): 317-320, 1991.
EA	EF	Ma, X-K. et al., "Carvedilol, a New Beta Adrenoreceptor Blocker and Free Radical Scavenger, Attenuates Myocardial Ischemia-Reperfusion Injury in Hypercholesterolemic Rabbits," <i>The Journal of Pharmacology and Experimental Therapeutics</i> 277(1): 128-136, April 1996.
EA	EG	Martin, J. et al., "Investigation on the new free radical scavenger polynitroxyl-albumin to prevent ischemia and reperfusion injury after orthotopic heart transplantation in the pig model," <i>European Journal of Cardio-thoracic Surgery</i> 19: 321-325, 2001.
EA	EH	Mitsos, S.E. et al., "Canine Myocardial Reperfusion Injury: Protection by a Free Radical Scavenger, N-2-Mercaptopropionyl Glycine," <i>Journal of Cardiovascular Pharmacology</i> 8(5): 978-988, September/October 1986.
EA	EI	Muldoon, L.L. et al., "Delayed Administration of Sodium Thiosulfate in Animal Models Reduces Platinum Ototoxicity without Reduction of Antitumor Activity," <i>Clinical Cancer Research</i> 6: 300-315, January 2000.
EA	EJ	Muldoon, L.L. et al., "Rescue from Enhanced Alkylator-Induced Cell Death with Low Molecular Weight Sulfur-Containing Chemoprotectants," <i>The Journal of Pharmacology and Experimental Therapeutics</i> 296(3): 797-805, 2001.
EA	EK	Nakano, H. et al., "Amelioration of Hepatocellular Integrity and Inhibition of Sinusoidal Oxidative Stress by N-Acetylcysteine Pretreatment in Cold Ischemia-Reperfusion Injury of Rat Liver," <i>European Surgical Research</i> 28: 245-255, 1996.

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FA						

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				YES	NO
FB					

OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

EA	FC	Neuwalt, E.A. et al., "Therapeutic Efficacy of Aortic Administration of N-Acetylcysteine as a Chemoprotectant against Bone Marrow Toxicity after Intracarotid Administration of Alkylators, with or without Glutathione Depletion in a Rat Model," <i>Cancer Research</i> 61: 7868-7874, November 1, 2001.
EA	FD	Neuwelt, E.A. et al., "First Evidence of Otoprotection Against Carboplatin-Induced Hearing Loss with a Two-Compartment System in Patients with Central Nervous System Malignancy Using Sodium Thiosulfate," <i>The Journal of Pharmacology and Experimental Therapeutics</i> 286(1): 77-84, 1998.
EA	FE	Neuwelt, E.A. et al., "In Vitro and Animal Studies of Sodium Thiosulfate as a Potential Chemoprotectant against Carboplatin-induced Ototoxicity," <i>Cancer Research</i> 56(4): 706-709, February 15, 1996.
EA	FF	No Author, "N-acetylcysteine," <i>Alternative Medicine Review</i> 5(5): 467-471, 2000.
EA	FG	Olanders, K. et al., "Protective Effects of N-Acetyl-L-Cysteine and Platelet Activating Factor Inhibition Are Not Linked to Intercellular Adhesion Molecule-1 Expression after Intestinal Ischemia and Reperfusion Injury in Rats," <i>Scandinavian Journal of Gastroenterology</i> 38: 618-625, 2003.
EA	FH	Oredsson, S. et al., "Allopurinol - A Free Radical Scavenger - Reduces Reperfusion Injury in Skeletal Muscle," <i>European Journal of Vascular Surgery</i> 5: 47-52, February 1991.
EA	FI	Pardridge, W.M., "Brain Drug Targeting and Gene Technologies," <i>Jpn J. Pharmacol.</i> 87: 97-103, 2001.
EA	FJ	Pugsley, W. et al., "The Impact of Microemboli During Cardiopulmonary Bypass on Neuropsychological Functioning," <i>Stroke</i> 25(7): 1393-1399, July 1994.
EA	FK	Qui, Y. et al., "The Influence of N-Acetylcysteine on Cardiac Function and Rhythm Disorders During Ischemia Reperfusion," <i>Cardioscience</i> 1(1): 65-74, March 1990.
EA	FL	Remsen, L. et al., "The Influence of Anesthetic Choice, Paco ₂ , and Other Factors on Osmotic Blood-Brain Barrier Disruption in Rats with Brain Tumor Xenografts," <i>Anesth. Analg.</i> 88: 559-567, 1999.

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	GB					
OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)						
EA	GC	Roach, G.W. et al., "Adverse Cerebral Outcomes After Coronary Bypass Surgery," <i>The New England Journal of Medicine</i> 335(25): 1857-1863, December 19, 1996.				
EA	GD	Salom, M.G. et al., "Protective effect of N-acetyl-L-cysteine on the renal failure induced by inferior vena cava occlusion," <i>Transplantation</i> 65(10): 1315-1321, May 27, 1998.				
EA	GE	Sehirli, A.O. et al., "Protective effect of N-acetylcysteine on renal ischemia/reperfusion injury in the rat," <i>Journal of Nephrology</i> 16: 75-80, 2003.				
EA	GF	Sekhon, B. et al., "N-Acetyl cysteine protects against injury in a rat model of focal cerebral ischemia," <i>Brain Research</i> 971: 1-8, 2003.				
EA	GG	Sekhon, C.S. et al., "Attenuation of renal ischemia/reperfusion injury by a triple drug combination therapy," <i>Journal of Nephrology</i> 16: 63-74, 2003.				
EA	GH	Sener, G. et al., "Melatonin and N-acetylcysteine have beneficial effects during hepatic ischemia and reperfusion," <i>Life Sciences</i> 72: 2707-2718, 2003.				
EA	GI	Shattuck, K.E. et al., "N-Acetylcysteine Protects From Glutathione Depletion in Rats Exposed to Hyperoxia," <i>Journal of Parenteral and Enteral Nutrition</i> 22(4): 228-233, July/August 1998.				
EA	GJ	Silbergleit, R. et al., "Lack of a neuroprotective effect from N-acetylcysteine after cardiac arrest and resuscitation in a canine model," <i>Resuscitation</i> 40: 181-186, 1999.				
EA	GK	Sochman, J. et al., "Cardioprotective effects of N-acetylcysteine: the reduction in the extent of infarction and occurrence of reperfusion arrhythmias in the dog," <i>International Journal of Cardiology</i> 28: 191-196, August 1990.				
EA	GL	Sochman, J., "N-Acetylcysteine in Acute Cardiology: 10Years Later. What Do We Know and What Would We Like to Know?!" <i>Journal of the American College of Cardiology</i> 39(9): 1422-1428, May 1, 2002.				
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HB					

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EA	HC	Tada, H. et al., "Effects of Pretreatment with 2-O-Octadecylascorbic Acid, a Novel Free Radical Scavenger, on Reperfusion-Induced Arrhythmias in Isolated Perfused Rat Hearts," <i>Journal of Cardiovascular Pharmacology</i> 16(6): 984-991, December 1990.
EA	HD	Tamura, Y. et al., "Reperfusion Injury in Bone: Effects of CV-3611, a Free Radical Scavenger, on Ischemic Revascularized Bone Grafts in Rats," <i>Journal of Reconstructive Microsurgery</i> 8(6): 471-479, November 1992.
EA	HE	Tan, D. et al., "Chemical and Physical Properties and Potential Mechanisms: Melatonin as a Broad Spectrum Antioxidant and Free Radical Scavenger," <i>Current Topics in Medicinal Chemistry</i> 2: 181-197, 2002.
EA	HF	Tang and Tang, "Protective Effects of SH-Compounds on Ischemia Reperfusion Induced Arrhythmias in the Isolation Rat Heart," <i>Acta Pharmaceutica Sinica</i> 26(2): 91-95, 1991.
EA	HG	Tepel, M. et al., "The Antioxidant Acetylcysteine Reduces Cardiovascular Events in Patients With End-Stage Renal Failure. A Randomized, Controlled Trial," <i>Circulation</i> 107: 992-995, February 25, 2003.
EA	HH	Thies, J.C. et al., "The efficacy of N-acetylcysteine as a hepatoprotective agent in liver transplantation," <i>Transplant International</i> 11(Suppl. 1): S390-S392, 1998.
EA	HI	Tian, H. et al., "Antioxidant N-acetylcysteine and AMPA/KA receptor antagonist DNQX inhibited mixed lineage kinase-3 activation following cerebral ischemia in rat hippocampus," <i>Neuroscience Research</i> 47: 47-53, 2003
EA	HJ	Tredger, J.M., "N-Acetylcysteine: Not Simply a Glutathione Precursor," <i>Transplantation</i> 69(5): 703-704, March 15, 2000.
EA	HK	Tripathi and Hegde, "Effect of N-acetylcysteine on myocardial infarct size following ischemia and reperfusion in dogs," <i>Indian J. Physiol. Pharmacol.</i> 42(1): 50-56, 1998.
EA	HL	Tsuji and Tamai, "Carrier-mediated or specialized transport of drugs across the blood-brain barrier," <i>Advanced Drug Delivery Reviews</i> 36: 277-290, 1999.

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EA	IC	van Jaarsveld, H. et al., "Effect of Normothermic Ischemic Cardiac Arrest and of Reperfusion on the Free Oxygen Radical Scavenger Enzymes and Xanthine Oxidase (a Generator of Superoxide Anions)," <i>Enzyme</i> 39: 8-16, 1988.				
EA	ID	Vanella, A. et al., "Free Radical Scavenger Depletion in Post-Ischemic Reperfusion Brain Damage," <i>Neurochemical Research</i> 18(12): 1337-1340, July 1993.				
EA	IE	Walcher, F. et al., "N-Acetylcysteine failed to improve early microcirculatory alterations of the rat liver after transplantation," <i>Transplant International</i> 8: 317-323, 1995.				
EA	IF	Weinbroum, A.A. et al., "Liver glutathione level influences myocardial reperfusion injury following liver ischemia-reperfusion," <i>Med. Sci. Monit.</i> 7(6): 1137-1144, 2001.				
EA	IG	Weinbroum, A.A. et al., "Lung preconditioning with N-acetyl-L-cystein prevents reperfusion injury after liver no flow-reflow: a dose-response study," <i>Transplantation</i> 71(2): 300-306, January 27, 2001.				
EA	IH	Weinbroum, A.A. et al., "N-Acetyl-L-Cysteine for Preventing Lung Reperfusion Injury After Liver Ischemia-Reperfusion," <i>Transplantation</i> 69(5): 853-859, March 15, 2000.				
EA	II	Xiong, Y. et al., "Effect of N-Acetylcysteine on Mitochondrial Function Following Traumatic Brain Injury in Rats," <i>Journal of Neurotrauma</i> 16(11):1067-1082, November 1999.				
EA	IJ	Yan and Greene, "Prevention of PC12 Cell Death by N-Acetylcysteine Requires Activation of the Ras Pathway," <i>The Journal of Neuroscience</i> 18(11): 4042-4049, June 1, 1998.				
EA	IK	Yip, L. et al., "Intravenous administration of oral N-acetylcysteine," <i>Critical Care Medicine</i> 26(1): 40-43, January 1998.				
EA	IL	Ytrebø, L.M. et al., "N-acetylcysteine increases cerebral perfusion pressure in pigs with fulminant hepatic failure," <i>Crit. Care Med.</i> 29(10): 1989-1995, 2001.				
EXAMINER			/Ernst Arnold/		DATE CONSIDERED 07/21/2006	
* EXAMINER: Initial if reference considered, whether or not criteria is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant(s).						